Atty. Docket No.: 015389-002640US

Docker No.: 013335-06204035

pplicant: Thomas R. Cech et al.

ERASE CATALYTIC SUBUNIT: DIAGNOSTIC AN

THERAPEUTIC METHODS

Sheet 1 of 34

1/34

Motif 0

ISEIEWLVLGKRSNAKMCLSDFEKRKQIFAEFIYWLYNSFIIPILQSFFYITESSDLRNR LKDFRWLFISD---IWFTKHNFENLNQLAICFISWLFRQLIPKIIQTFFYCTEISSTVT-TREISWMQVET-SAKHFYYFDHEN-IYVLWKLLRWIFEDLVVSLIRCFFYVTEQQKSYSK AKFLHWLMSVYVVELLRSFFYVTETTFQKNR

human

tez1 EST2 p123

human

EST2

p123

tez1

Motif 1

TVYFRKDIWKLLCRPFI-TSMKMEAFEKINENNVRMDTQK-TTLPPAVIRLLPKK--NTF IVYFRHDTWNKLITPFIVEYFKTYLVENNVCRNHNSYTLS--NFNHSKMRIIPKKSNNEF TYYYRKNIWDVIMKMSI-ADLKKETLAEVQEKEVEEWKKS-LGFAPGKLRLIPKK--TTF LFFYRKSVWSKLQSIGIRQHLKRVQLRDVSEAEVRQHREARPALLTSRLRFIPKP--DGL

human

EST2 5123

tez1

RIIAIPCRGADEEEFTIYKENHKNAIQPTQKILEYLRNKRPTSFTKIYSPTQIADRIKEF RLITN-LRKRFLIKMGSNKKMLVSTNQTLRPVASILKHLINEESSGIPFNLEVYMKLLTF RPIMTFNKKIVNSDRKTTKLTTNTKLLNSHLMLKTLKN-RMFKDPFGFAVFNYDDVMKKY RPIVNMDYVVGARTFRREKRAERLTSRVKALF-SVLNYERA

tez1 EST2

p123

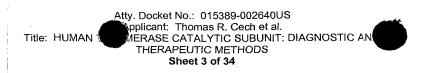
KKDLLKHRMFGR-KKYFVRIDIKSCYDRIKQDLMFRIVKK-KLKDPEFVIRKYATIHATS KQRLLKKFINIVLPELYFMKFDVKSCYDSIPRMECMRILKD-ALKNENGFFVRSQYFFNTN EEFVCKWKQVGQPKLFFATMDIEKCYDSVNREKLSTFLKTTKLLSSDFWIMTAQILKRKN Motif 3 (A)

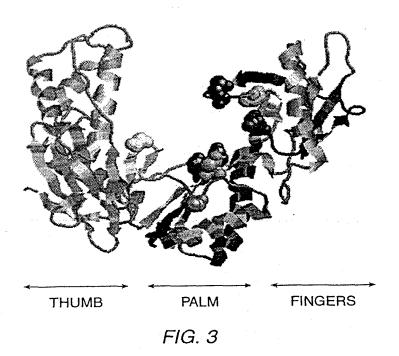
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Applicant: Thomas R. Cech et al.

MERASE CATALYTIC SUBUNIT: DIAGNOSTIC AN
THERAPEUTIC METHODS
Sheet 2 of 34

2/34

ht pl 10.6 11.3 10.1		
Mol. weight 116.000 127.000 123.000 103.000		
Sp_Trt1p - hTRT - Ea_p123 Sc_Est2p		
T 12 A B'CD E	EMPZZ(二)-(Z)-(B) (B)	and (Copia-Ty1) and ZZACTACTACTACTACTACTACTACTACTACTACTACTACTA
	msDNAs Mito.plasmid/RTL Group II introns Non-LTR Retrotransposons	Hepadnaviruses LTR Retrotransposons (Copia-Ty1) LTR Retrotransposons (Gypsy-Ty3) Caulimoviruses Retroviruses
50 aa		





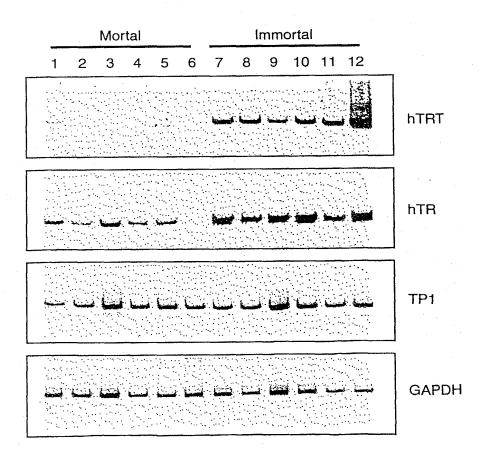


FIG. 5

32

ga flu

ga äin i isapis

Motif T

TRT con Sp_Trt1p hrrr Ea_p123 Sc_Est2p	WL hh hh p ffy te p p y rk w l h i k 429 wlynsfiipilosf fytte sdlrnrtvyfrkdiwkllcrpfitsmkm 8 546 wlmsvyvvellrsffyrnrlffyrksvwsklosigirqhlk 10 441 wifedlvvslircffyvtgrsktyyrkniwdvimkmsiadlkk 8 366 wlfrolipkiiqtffycteisstvt-ivyfrhdtwnklitpfiveyfr 8
TRT con Sp_Trt1p hTRT Ea_p123 Sc_Est2p	Motif 1 hhk K helipek p Frhi h h relyfhhdh CYD i hhk K hnvermotorteppavirleppa
RT con Sc_a1 Dm_TART HIV-1	D hh h K LSNELGTGKFKFKFRPMRIVNI PK PKGG 0 IRPLSVGNPRDKIVQEVMRMILDTIFDKK 27 FGGSNW F IEV D LKK CFDTI SHDLIIKEL K RYISD 20 S SILRIGYYPDAWKHAQVKMILKPGKS 6 YRPISLLSGLSKMFERLLLKRLFRVDLFK 32 RKEYCSAVFL D ISEAF D RVWHEGLLLKLAKILPY 25 (EGKISKIGPENPYNTPVFAIKKKDST 1 WRKLVDFRELNKRTQDFWEVQLGIPHPAG 0 LKKKKSVTVL D VGDAYFSVPLDEDFRKYTAFTIP 7
TRT con Sp_Trt1p hTRT Ea_p123 Sc_Est2p	TRT con K Y Q GIPQGS LS hL h Y DL F LLRL DDFLhIT A F h G c p N cK W G S **ACTIVE CON K Y Q GIPQGS LS hL h Y DL F LLRUVDDFLFITVNKKD 0 AKKFLNLSLRGFEKHNFSTSLEKTVI 17 KKRMPFFGFSV 181 **ACTIVE CON K Y Q GIPQGS LS HL H H L LARVVDDFLFITVNKKD 0 AKKFLNLSLRGFEKHNFSTSLEKTVI 17 KKRMPFFGFSV 181 **ACTIVE CONTROL

55 YVRYADDILIGVLGSKN 2 KMIKRDLNNFLNS-LGLTMNEEKTLI 4 ETPARFLGYNI 7 LSTYADDTIVLSSDILA 6 NENYLKTFSDWADKWGISVNAAKTGH 25 ESKQSYLGVIL 4 IYQYMDDLYVGSDLEIG 1 HRTKIEELRQHLLRWGLTTPDKKHQK 0 EPPFLWMGITL

c**K**

Gh h

Y DDhhh

Ч

RAGOIGA**GVPOGSNLGPILYSIFSSDMPLPHIYHP** GIRY**Q**YNVL**PQGW**KG**S**PAIFQSSMTKILEPFKKON hpgg pP hh h TYHKPMLGLPQGSLISPILCNIVMTLVDNWLEDYI

Sc_a1 Dm_TART HIV-1

RT con

Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
Title: HUMAN DMERASE CATALYTIC SUBUNIT: DIAGNOSTIC A
THERAPEUTIC METHODS
Sheet 5 of 34

5/34

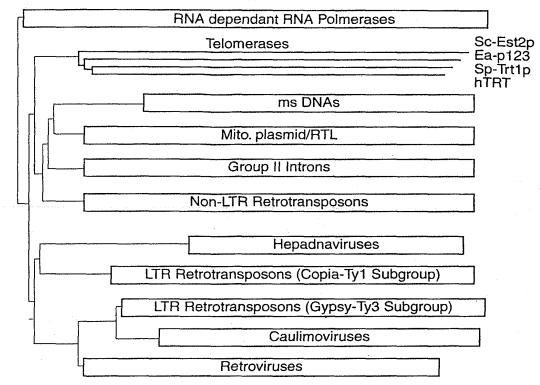


FIG. 6

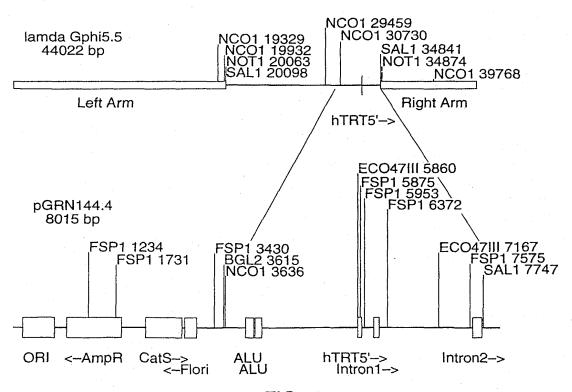


FIG. 7

an in

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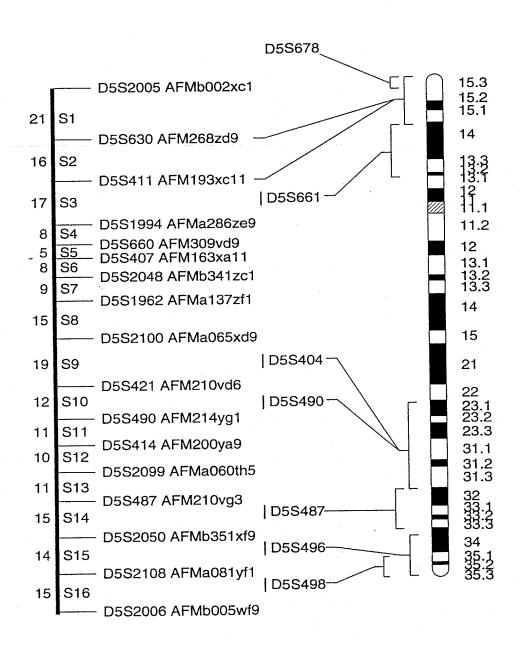


FIG. 8

7/34

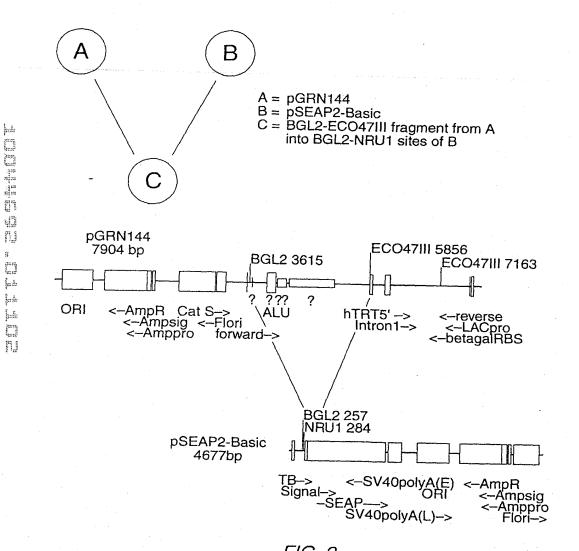


FIG. 9

Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
MERASE CATALYTIC SUBUNIT: DIAGNOSTIC AN
THERAPEUTIC METHODS
Sheet 8 of 34

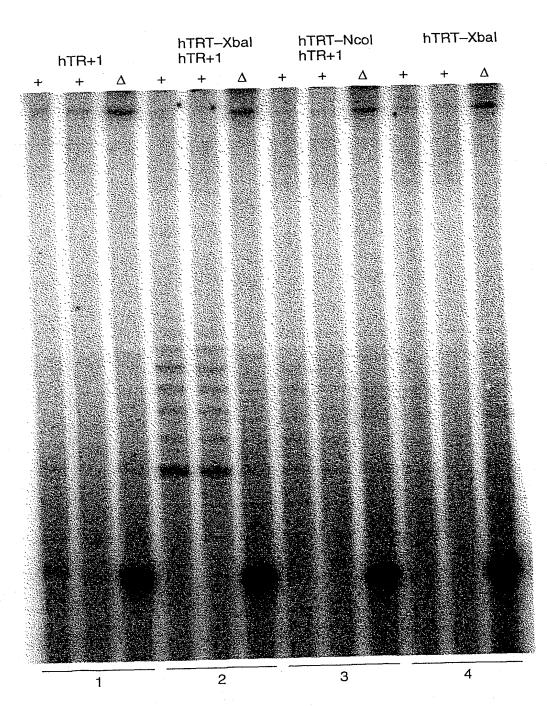


FIG. 10A

Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
MERASE CATALYTIC SUBUNIT: DIAGNOSTIC AI
THERAPEUTIC METHODS
Sheet 9 of 34

9/34

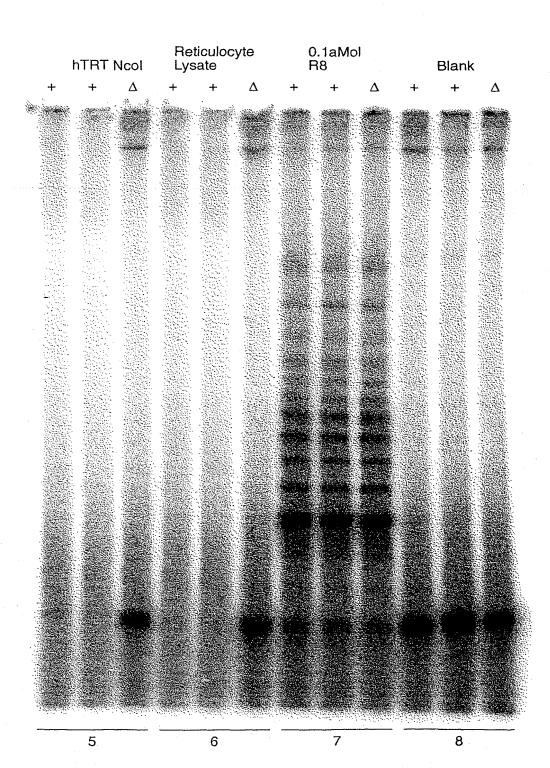


FIG. 10B

Atty. Docket No.: 015389-002640US
Doplicant: Thomas R. Cech et al.
Title: HUMAN TEMPERASE CATALYTIC SUBUNIT: DIAGNOSTIC AND
THERAPEUTIC METHODS
Sheet 10 of 34

Telomerase Specific Motifs

MOTIF T'	EV	EAEVR	ENNVR	EKEVE	9 ENNVC
		13	12	12	
	Η	IGI	PFI	MSI	PFI
	y Rk W l	KSVWSKLQS:	KDIWKLLCR!	KNIWDVIMK	HDTWNKLIT
	α ≻	FYR	/YFR	YYYB	NFR
MOTIF T	Wl FFY TE	6 WLMSVYVVELLRSFFYVTETTFQKNRLFFYRKSVWSKLQSIGI 13	9 WLYNSFIIPILQSFFYITESSDLRNRTVYFRKDIWKLLCRPFI	1 WIFEDLVVSLIRCFFYVTEQQKSYSKTYYYRKNIWDVIMKMSI	6 WLFRQLIPKIIQTFFYCTEISSTVT.IVYFRHDTWNKLITPFI
		546	429	44	366
	TRT con	hTRT	spTRT	Ea_p123	Sc_Est2

Telomerase RT Motifs (Fingers)

MOTIF B'	Y q GipQGs 1S 1 y	104		PKLFFATMDIEKCYDSV 117 YKQTKGIPQGLCVSSILSSFYY	85 YIREDGLFQGSSLSAPIVDLVY		
MOTIF A	p lyF D cYD i	69 PELYFVKVDVTGAYDTI	66 RKKYFVRIDIKSCYDRI	67 PKLFFATMDIEKCYDSV	68 PELYFMKFDVKSCYDSI	h hDh AF h	λS
MOTIF 2	fr I	0 LRPIV	0 FRLIT	O FRPIM	2 FRIIA	hR h	
MOTIF 1	R iPKk	SRLRFIPKPDG 0 LRPIV	AVIRLLPKKNT 0 FRLIT	GKLRLIPKKTT 0 FRPIM	SKMRIIPKKSN 2 FRIIA	p hh h K	
		근	10	10	13		
	TRT con	hTRT	spTRT	Ea_p123	Sc_Est2	RT con	

Telomerase RT Motifs (Palm, Primer Grip)

		192	176	174	141		
MOTIF E	wgs l	WCGLLLDTRTL	FFGFSVNMRSL	15 VSRENGFKFNMKKLQT 28 WIGISIDMKTL	WKHSSTMNNFH	hLG h	
		24	22	28	25		
		2	IΛ	Ŋ	LA	ᅜ	
_	×	JRKJ	EKJ	KKI	Č	Gh h cK h	
<u>H</u>	n R	MNI	STSI	(FIN	SANI	ᅺ	
MOTIF D		(GC)	INF	IGFF	NA	Сh	
4		LLLRLVDDFLLVT 15 GVPEYGCVVNLRKTVV 24	16 VLLRVVDDFLFIT 15 GFEKHNFSTSLEKTVI	SRE	LILKLADDFLIIS 15 GFQKYNAKANRDKILA 25		
	b	5	ις. O	5	S		
	بد	ĬŢ.	E	1.1	S.		
ິບ	111rl DDfL it	LIL	LFI	LLMRLTDDYLLIT	LII	hh	
MOTIF C	B	7DDF	7DDE	ď	DDE	и у орььь	
MO	디	RLV	RV	RL1	KLA	×	2
	111	LI	VLI	LIN	LI	Ч	
		15	16	24	18		
٠.	uc			53	£2	٦.	
	TRT con	RT.	SPTRT	Ea_p123	Sc_Est2	CO	
	TR	hTRT	gs	Ба	သို	RT	

Intron1

301 GCTGGGGTTGAGGGCGGCGGGGGAACCAGCGACATGCGGAGAGCAGCGCAGCGACTCCGACCCCAACTCCCGCCGGCCCCCTTGGTCGCTGTACGCCTCTCGTCGCGTCCGCTGAG

NFkB_CS1 GGGRQTYYQC NFkB_CS2 RGGGRMTYYCC

Topo_II_cleavage_site RNYNNCNNGYNGKTNYNY *******

361 AGGGCGCTTCCCCCGCAGGTGTCCTGCCTGAAGGAGCTGGTGGCCCGAGTGCTGCAGAGG TCCCGCGAAGGGGGCGTCCACAGGACGGACTTCCTCGACCACCGGGCTCACGACGTCTCC

1	AAAACCCCAA			AGCCCTGCAG		
51	AACCTCAGTA	TTAATAAGCT	CAGATTTTAA	ATATTAATTA	CAAAACCTAA	
101	ATGGAGGTTG	ATGTTGATAA	TCAAGCTGAT	AATCATGGCA	TTCACTCAGC	
151	TCTTAAGACT	TGTGAAGAAA	TTAAAGAAGC	TAAAACGTTG	TACTCTTGGA	
201	TCCAGAAAGT	TATTAGATGA	AGAAATCAAT	CTCAAAGTCA	TTATAAAGAT	
251	TTAGAAGATA	TTAAAATATT	TGCGCAGACA	AATATTGTTG	CTACTCCACG	
301	AGACTATAAT	GAAGAAGATT	TTAAAGTTAT	TGCAAGAAAA	GAAGTATTTT	
351	CAACTGGACT	AATGATCGAA	CTTATTGACA	AATGCTTAGT	TGAACTTCTT	
401	TCATCAAGCG	ATGTTTCAGA	TAGACAAAAA	CTTCAATGAT	TTGGATTTCA	
451	ACTTAAGGGA	AATCAATTAG	CAAAGACCCA	TTTATTAACA	GCTCTTTCAA	
501	CTCAAAAGCA	GTATTTCTTT	CAAGACGAAT	GGAACCAAGT	TAGAGCAATG	
551		AGCTCTTCCG		ACTAAATATT		
601	and the second of the second o	GAAGGAACTC	the second secon	TTGCGGGAAT	AACGTTTTTG	
651				AAAAGCAAAA		
701				ACCTGCAAAT		
751		GATCACTTTC		CAACGTGCCG		
801				GCACTCATTT		
851	AACCAATTCT			AGTAACAAAA		
901	AGCGATGGAC			GAATATATTC		
951				TCGAAAAAAT		
1001				TACTATTTAA		
1051				ACAAAAAATC		
1101	TAAATAAAAC			ACTATGAAGA		
1151				TTTATTAATG		
1201	CAATATACTC			AAGAAACCGT		
1251	AAAAGAAAGT			ACAAGCATGA		
1301	AAAAACTTAT			AGAGAAATAT		
1351	GGTTGAGACC			TTTTGATCAC		
1401				TCGAGGATCT		
1451				CAACAGAAAA		
1501	AACCTATTAC			CGTCATTATG		
1551				AGGTCCAAGA		
				CCTGGAAAAC		
1601						
1651				GACTTTCAAT		
1701				CTACAAATAC		
1751	AACTCTCACT			AATAGAATGT		
1801	TTTTGGATTC			TGTAATGAAA		
1851		CAAATGGAAG		AACCAAAACT		
1901	ACTATGGATA			GTAAACAGAG		
1951		AAAACTACTA		TTCAGATTTC		
2001	CTGCACAAAT	-	AAGAATAACA	TAGTTATCGA		
2051		AAGAAATGAA		AGACAGAAAT		
2101				ATTCAGTGTT		
2151		CTTAAATGCA		TAATTGTTGA		
2201	AGAAATTATT			CAACCAGTCA		
2251	CCAATATAAT					
2301				TTTTGTCATC		
2351	GCAACATTAG	AGGAAAGCTC	CTTAGGATTC	CTTAGAGATG	AATCAATGAA	

2401	CCCTGAAAAT	CCAAATGTTA	ATCTTCTAAT	GAGACTTACA	GATGACTATC
2451	TTTTGATTAC	AACTCAAGAG	AATAATGCAG	TATTGTTTAT	TGAGAAACTT
2501	ATAAACGTAA	GTCGTGAAAA	TGGATTTAAA	TTCAATATGA	AGAAACTACA
2551	GACTAGTTTT	CCATTAAGTC	CAAGCAAATT	TGCAAAATAC	GGAATGGATA
2601	GTGTTGAGGA	GCAAAATATT	GTTCAAGATT	ACTGCGATTG	GATTGGCATC
2651	TCAATTGATA	TGAAAACTCT	TGCTTTAATG	CCAAATATTA	ACTTGAGAAT
2701	AGAAGGAATT	CTGTGTACAC	TCAATCTAAA	CATGCAAACA	AAGAAAGCAT
2751	CAATGTGGCT	CAAGAAGAAA	CTAAAGTCGT	TTTTAATGAA	TAACATTACC
2801	CATTATTTTA	GAAAGACGAT	TACAACCGAA	GACTTTGCGA	ATAAAACTCT
2851	CAACAAGTTA	TTTATATCAG	GCGGTTACAA	ATACATGCAA	TGAGCCAAAG
2901	AATACAAGGA	CCACTTTAAG	AAGAACTTAG	CTATGAGCAG	TATGATCGAC
2951	TTAGAGGTAT	CTAAAATTAT	ATACTCTGTA	ACCAGAGCAT	TCTTTAAATA
3001	CCTTGTGTGC	AATATTAAGG	ATACAATTTT	TGGAGAGGAG	CATTATCCAG
3051	ACTTTTTCCT	TAGCACACTG	AAGCACTTTA	TTGAAATATT	CAGCACAAAA
3101	AAGTACATTT	TCAACAGAGT	TTGCATGATC	CTCAAGGCAA	AAGAAGCAAA
3151	GCTAAAAAGT	GACCAATGTC	AATCTCTAAT	TCAATATGAT	GCATAGTCGA
3201	CTATTCTAAC	TTATTTTGGA	AAGTTAATTT	TCAATTTTTG	TCTTATATAC
3251_	TGGGGTTTTG	GGGTTTTGGG	GTTTTGGGG		

FIG. 13 (CONTINUED)

	1	MEVDVDNQAD	NHGIHSALKT	CEEIKEAKTL	YSWIQKVIRC	RNQSQSHYKD
	51	LEDIKIFAQT	NIVATPRDYN	EEDFKVIARK	EVFSTGLMIE	LIDKCLVELL
	101	SSSDVSDRQK	LQCFGFQLKG	NQLAKTHLLT	ALSTQKQYFF	QDEWNQVRAM
	151	IGNELFRHLY	TKYLIFQRTS	EGTLVQFCGN	NVFDHLKVND	KFDKKQKGGA
	201	ADMNEPRCCS	TCKYNVKNEK	DHFLNNINVP	NWNNMKSRTR	IFYCTHFNRN
	251	NOFFKKHEFV	SNKNNISAMD	RAQTIFTNIF	RFNRIRKKLK	DKVIEKIAYM
	301	LEKVKDFNFN	YYLTKSCPLP	ENWRERKQKI	ENLINKTREE	KSKYYEELFS
	351	YTTDNKCVTQ	FINEFFYNIL	PKDFLTGRNR	KNFQKKVKKY	VELNKHELIH
٠.,	401	KNLLLEKINT	REISWMQVET	SAKHFYYFDH	ENIYVLWKLL	RWIFEDLVVS
	451	LIRCFFYVTE	QQKSYSKTYY	YRKNIWDVIM	KMSIADLKKE	TLAEVQEKEV
	501	EEWKKSLGFA	PGKLRLIPKK	TTFRPIMTFN	KKIVNSDRKT	TKLTTNTKLL
	551	NSHLMLKTLK	NRMFKDPFGF	AVFNYDDVMK	KYEEFVCKWK	QVGQPKLFFA
	601	TMDIEKCYDS	VNREKLSTFL	KTTKLLSSDF	WIMTAQILKR	KNNIVIDSKN
	651	FRKKEMKDYF	RQKFQKIALE	GGQYPTLFSV	LENEQNDLNA	KKTLIVEAKQ
•	701	RNYFKKDNLL	QPVINICQYN	YINFNGKFYK	QTKGIPQGLC	VSSILSSFYY
•	751	ATLEESSLGF	LRDESMNPEN	PNVNLLMRLT	DDYLLITTQE	NNAVLFIEKL
	801	INVSRENGFK	FNMKKLQTSF	PLSPSKFAKY	GMDSVEEQNI	VQDYCDWIGI
	851	SIDMKTLALM	PNINLRIEGI	LCTLNLNMQT	KKASMWLKKK	LKSFLMNNIT
	901	HYFRKTITTE	DFANKTLNKL	FISGGYKYMQ	CAKEYKDHFK	KNLAMSSMID
	951	LEVSKIIYSV	TRAFFKYLVC	NIKDTIFGEE	HYPDFFLSTL	KHFIEIFSTK
1	001	KYIFNRVCMI	LKAKEAKLKS	DOCOSLIOYD	A	

000000000	8	8	88	8	2	225	05	65 ×
80 1460 2240 320 4400 640 720 880 958	1018 20	1078 40	1138 60	1198 80	1272 86	1332 106	14	1469 128
ggtaccgatttactttcctttcttcataagctaattgcttcctcgaacgctcctaaatctctggaaatatttttacaaga acteaataaagaacaattccaatatgaaggtgttattagtgatcgataattttctattttattttatttctattttcattttcattttcg ccaagtataaagaccaaacattccttccccctaaagacttttacttttattatttctattttcaatatttcg ggttcgcttacttttaatcgtggtactgttttagctgctacttctagccaaccgcgttttctaacccgtcattggatat agctcttggagtagctcacagaaatccttacaaatcttctgatgagaccaaccgcgtgtttcatcacaagccgttggatattcttaacaggagccttacacagaaatccttacaaatcttctgatgagaccatattctagagagccttacacttagagggccgtgggaaagctcttgggggaaagttttggtatcatccaacgtttggcttggaaagggttgataaaaaaattctaaaaggagccttacacttgggggaaacccttagagggaaagccttaacagccgtttggggaaaggttgggaaaggttggtatcatttttctattttctatttctattgttgttccaaaaattgagatagagaaacccttttaccactagagaaagattggtaaccgtccttttaattttcatttttaatggtttaaaattgataaaaaaaa	GTA V	TCG S	CTT L	CCA P	gtatatattttttgttttgattttttttctattcgggatagctaatatgggcag	CTA	gtaaggtattctaattgtgaaatatttacctgcaattactgtttcaaagaga	AAT N
ttac atatac intega intega intega intega intega intega intega intega	TAT Y	AGC S	$ ext{TTT}$	TCT	atgg	AAT N	caaa	CAA Q
gatttactttccttcttcataagctaattgcttcctcgaacgctcctaaatctctggaaatatttttacaag taacaataccaagtcaaattccaatagaaggtgttattagtgatcgataatatttctatttatctggtcgt. ataaggacaaaaagaacaacttccttccccctaaagacttttactttattta	CAA	GCA	ATT I	TCT	latat	AGG AAT C R N I	gtt	CGA GCC ATG CAT GTA AAC GGA GTA CAA AAT R A M H V N G V Q N
yaaat tatttc tttcccccccccccccccccccccccc	AAT N	CCG P	TCT S	TTT F	igote	CGA R	tact	GGA G
tetogen construction of the construction of th	GAG E	TCG S	TTT	CAA	ggatë	CGT R	ycaat	AAC N
satct taatt ytgtt ytgtt yattc yattcat ttgat ccttt actct actcc	CCC AAA AGC AGG ATT CTT CGC TTT CTAP F K S R I L R F L	999 9	TCC	GTT V	ttag	AAA CAG ATG TTC GAT GAA AGT TTT GAG CGT CGA K Q M F D E S F E R R	accto	GTA V
cottage cottag	TTT F	AGA R	ACG T	GGT G	tctat	TTT F	attt	CAT H
cgct tract ccaa ccaa ccaa ccaa ccaa ccaa c	CGC R	$ ext{TTG}$	CAA Q	GAA	ttti	AGT S	aaat	ATG M
cgaa cttagaa ccaagaa ccaagaa ccaagaa ccaagaa ccaagaa ccaagaa catta	CTT	GTT V	GTA V	GAT	gatt	GAA	tgtg	GCC
tcct garta aactt taat taat taat taat taat aaaa aaaaa	ATT	CTT L	GAT D	CCA P	tt	GAT	taat	CGA R
tgcttcctcctcctcctccccccccccccccccccccc	AGG R	CAA Q	AGC S	AAG K	tttg	$ ext{T}^{ ext{T}}$	attc	GAA GAT TTT E D F
taat tocc tagc aaat aacgt aacgt agtag agtaa	AGC	GTA	AGA R	AGT	tatt	ATG	aggt	GAT D
aaagc ccaat tcct tcag cctag cctag tccat tcag	AAA K	$_{\rm Y}^{\rm TAT}$	TTG	GAC	tata	CAG Q		GAA
tcat tact tact tact tca tca tca t	CCC	GAT D	CGC	TTC	gta	AAA K	ATG M	CAT
tttcaag ggaaaa attaaaa attaaaaa attaaaa attaaaa	ACC	AAT N	GAA	0 0	GAG E	GTA V	TCC	AAT N
tcct a a a a a a c c a c a c a c c a c a a a c a c	CAT H	TTA L	TGC	GTC V	TCA	GTT V	TTT F	taaccgataaag
actt Gatact Catac Catact Catact Catact Catact Catact Catact Catact Catact Catact Catact Catact Catact Catact Catact Catac Catac Catac Catact Catac Cata Cata	CAC	ACC T	ATA	GTA V	CAG Q	AAT		cgat
attragad traca addaga traca adtra adtra attra attra	GAA	TGT	AAT N	ACT T	TCA S	GCG A	AAA K	taac
ugtaccg acteaata caagta ggttcgc ggttcgc ttgacat ttgaga ttgaga ttgaga ttgaga ttgaga ttgaga ttgaga ttgaga ttgatct	ACC	CTA L	AGC	TCG	$^{TGC}_{C}$	ATA I	ATG M	u
acteaal ccaagte ggttcgc agctctt ttaacat gttgaca attgage ccaaate ataatct gatactt	ATG M	TAC Y	TAT Y	CAT H	AAA K	CTA L	$_{\rm L}^{\rm cTG}$	ttgtat
181 224 324 440 721 721 880 881	959 1	1019 21	1079	1139	1199	1273 87	1333	1406 114

1529 148	1601 155	1661 175	1721 195	1781 215	1841 235	1907 245	1967 265	2027 285	2087 305	2147 325	2207 345	2267 365
CAA Q	3 3 9	GAC D	GTG V	AAA K	TAT Y	AAC N	AGG R	GTA V	ACA T	ATT I	GCG A	ATA I
TGG W	ATC	AAT N	ACT	CGC R	TCC	TTT AZ F N	CCA P	CTG L	CAA Q	TAT Y	TTT F	AGG R
AAT	ag T	CCA	GAA	GCC	TCA	TAT TY Y	TTT	CCA	GAA E	CCA	GTG	CAA Q
AAA K	gtaaataccggttaagatgttgcgcactttgaacaagactgacaagtatag	CTT	GAG E	AGC S	AGG R	CTA TI	ATT I	ATT I	ATT	TGC	CAG Q	AAC N
TCA S	acaa	GCT A	TTT F	AAA K	TAC Y	AT C	TGG W	GTG V	TTA	TAT Y	, AAC N	G GGT
GAG	actg	GAG E	GTG V	AAT N	TTT F	G gtaactaatactgttatccttcataactaattttag AT D	CAA Q	AAA K	CCT P	CAT	F CCG	TGG W
CTT L	срад	$\Gamma \Gamma \Gamma$	AAT N	CAA Q	ATT I	attti	CTT	CAC	TAC	AAC	AAG K	A ATC I
ATA I	tgaa	ATT I	AAT N	ACT T	AGC S	acta	TGG W	TTG	GTA V	TAC	TTA L	TTA L
TCT S	actt	AGT	AAA K	ATT I	<u> </u> ዮጥጥ F	cata	ATG M	CAA Q	AAG K	GTT V	TCC S	AAA K
ATA I	gcga	GGA	$ ext{TTT}$	TCC	AGG R	cctt	CAC H	AAG K	CTA	AAA	TAT Y	CCT P
CTT	tgtt	AAA K	$_{\rm L}^{\rm CTT}$	ACA	AGT	ctato	GTA V	GTG	crc	TCA	AGT S	TTT F
TAC Y	aaga	TCC	CCA P	GAA E	ATT	actg	ACA T	CAA	CGT R	CTA	CTT L	GTG V
AAT N	ggtt	TTA L	ATA I	ATT I	TCA S	caati	AAC N	TTT F	AAA K	TCT	, ATC I	, CGA
CCT	tacc	TTA L	399 9	ACC T	ATT I	caact	CGG R		CCC	ATT I	, AAA K	, GTT V
TTT F	taaa	TAC Y	TCT	CGA R	AGC S		GAT D		GTG V	CGT R	GAA	CTT
ACT	AT g I	CAT H	ATT I	AAG K	AAT N	CAA			GTT V	CAT	GAT	ATT
TCT	GAA E	ATG M	CAG Q	AGA R	TGG W		ATT I		ACA	CTC L	GAT	
GTT V		GCC	CTT	AAA K	TCC	TTT F	TCT	GGA G	AGT S			
CTC	ĪĞ	GAT	TAC Y	AAA K	${ m GTT}$	AAG K	CAC	TTT F	CAG Q		ACC	
GAT	TT	AGT	AT	TCA S	GAA E	AAG K	$ ext{TTA}$	CAA			GAC	
1470 (02					1908	1968 266	2028	2088	2148	04
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FIG. 15 (CONTINUED)

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2336 375	2396 395	2465 405	2525 425	2585 445	2645 465	2705 485	2775 495	2835 515	2906 524	2967 542	3027 562	3088 581
ACT T	AAG K	GGA G	GCG A	TAT Y	AAA K	GAG E	ACT T	ACG T	gtattaatttttggtcatcaatgtactttacttctaatctatta	GTG V	GAG E	gtaat
GAA E	ATA I	CTT L	TTT F	$ ext{TTT}$	TGG W	AAC N	AAA K	ATT I	atct	CCT P	$ ext{T}^{ ext{T}}$	უ ლ
CTC	AAC	GTC V	ATA	TTT	ATT I	ATA	cAG Q	CTC L	tcta	CGA R	AAC N	TTT F
y AC	AGT S	CTA (CAA	TCT	GAT D	AAA K	ACT	CGT R	tact	TTA L	TTT F	ATG M
accaç	ATG A	TGG (AAG K	CAA Q	AAA K	GAA E	GAT	TTT F	actt	ACT T	CCA	CGA R
G gtattgtataaaatttattaccactaacgattttaccag D	TTA 1	GAA S	CGC 7	TTA L	AGA R	TTT F	ATG	ACC	ıatgt	CAA Q	ATT I	CAC H
acgat	TAT Y	ATT (AAA K	ATT	TTT F	GCG A	AGG R	AAT N	atca	AAC	GGT G	AAG K
actaí	CAT ?	GAA	GAG	CCT	TAT Y	GAA	GTT V	AAG K	ggto	ACG T	AGT S	CTT
Tacci	TTA (L	TCA (TTT	ATA	GTT V	ATG M	AAT	AAG K	ttt	AGT	AGT S	CTT
tatt	AGT S	ATT I	GAT '	ATA	ACT	AAA K	AAC	CCT P	taat	GTC	GAA E	GAT
aaatt	TTT /		AGT (TTT	CGA R	ATG M	tcag	TTA L		TTA L	GAA E	AAG K
tatai	TCT S	caat	TTA	TCG S	AAT	TCA S	attt	CTA	AAG K	ATG M	AAT N	AAG K
attgi	GAG '	ttaa	TGC C	AAT	CGA. R	ACA	taat	CGT	ATA	AAA K	ATC I	TTT F
g gt.	TAC (Y	ttaa	ATG M	TAC Y	TTA L	ATT I	aago	ATT I	TTA L	AAA K	TTA L	ACT T
AAA C	AGA '	acca	AAA K	CTA	GAT	TTT F	caaa	GTT V	$ ext{TTC}$	AAC	CAT H	CTT
TTA 1	TCG Z	ccaaatttttttaccattaattaacaatcag	GCG A	TGG W	AGT S	CCC P	aagtattttttgcaaaaagctaatatttcag	GCA	AGA R	TCA S	AAA K	CTT
ATA '	TTA '	attt	AAT N	TAC	TCA S	CGA R	attt	CCA P	AAA K	GGT	CTG L	AAG K
ATA	AAA	ccaa	TCA	ATC I	GAA E	TGC C	laagt	CCT	AGA R	ATG M	ATA I	ATG M
GAG	TTG	. b	AGG R	$ ext{TTC}$	ACT T	TTG	ttta	TTG	TTA L	gcag	TCG	TAC
TTT (TTC ' F	gtaatat	AAA K	GAA E	ATC I	CTC	gtattt	ACT T	AAT N	ttag	GCA	GTT V
2268 366	2337	2397	2466	2526 426	2586 446	2646 466	2706	2776 496	2836 516	2907	2968	3028

FIG. 15 (CONTINUED)

17/34

3155 591	3215 611	3275 631	3343 643	3405 659	3465 679	3532 692	3593 708	3653 728	3713 748	3777 764	3840 778	3900 798
3089 tatataatgcgcgattcctcattatttacag G CGT AAG AAG TAT TTT GTA CGG ATA GAT ATA 31 582	3156 AAA TCC TGT TAT GAT CGA ATA AAG CAA GAT TTG ATG TTT CGG ATT GTT AAA AAG AAA CTC 32.592 K S C Y D R I K Q D L M F R I V K K L \sim 61.	216 AAG GAT CCC GAA TTT GTA ATT CGA AAG TAT GCA ACC ATA CAT GCA ACA AGT GAC CGA GCT 32'612 K D P E F V I R K Y A T I H A T S D R A 63	276 ACA AAA AAC TTT GTT AGT GAG GCG TTT TCC TAT T gtaagtttattttttttggaattttttaacaa 33	3344 attetttttag TT GAT ATG GTG CCT TTT GAA AAA GTC GTG CAG TTA CTT TCT ATG AAA ACA 34 644	3406 TCA GAT ACT TTG TTT GTT GAT TAT TGG ACC AAA AGT TCT TCT GAA ATT TTT 34. 660 S D T L F V D F V D Y W T K S S S E I F 67	3466 AAA ATG CTC AAG GAA CAT CTC TCT GGA CAC ATT GTT AAG gtataccaattgttgaattgtaataaca 35 680 K M L K E H L S G H I V K	3533 ctaatgaaactag ATA GGA AAT TCT CAA TAC CTT CAA AAA GTT GGT ATC CCT CAG GGC TCA 35 693 I P Q G S 70	3594 ATT CTG TCA TCT TTG TGT CAT TTC TAT ATG GAA GAT TTG ATT GAT GAA TAC CTA TCG 36 709 I L S S F L C H F Y M E D L I D E Y L S 72	3654 TTT ACG AAA AAG AAA GGA TCA GTG TTG TTA CGA GTA GTC GAC GAT TTC CTC TTT ATA ACA 37729 F T K K K G S V L L R V V D D F L F I T 74	3714 GTT AAT AAA AAA AAA TTT TTG AAT TTA TCT TTA AGA G GLGAGTLGCLGTCC 37 $^{\prime}$ 749 V N K K D A K K F L N L S L R G	3778 taagttotaaccgttgaag GA TTT GAG AAA CAC AAT TTT TCT ACG AGC CTG GAG AAA ACA GTA 38 765	3841 ATA AAC TTT GAA AAT AGT AAT GGG ATA ATA AAC AAT ACT TTT TTT AAT GAA AGC AAG AAA 39779 I N F E N S N G I I N N T F F N E S K K 79
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18/34

4209 888 4274 903 4468 946 4528 966 4588 986 4665 989 4339 4401 935 GCC A gtacgtgtc gtgagtacttatttaactaga tgtcattttcaatttattatatacatcctttattactggtgtcttaaaacaatattattactaagtata 999 AAA AAA K TTG AGA CAT ATG gtatactgtgtaactgaataatagctgacaaataatcag AAG K GAT GGT AGA AAA K AAA AAA K AAA K ACG AGA ATT I GAA ATG CTG AAT 0 A AAA K GAG GGA CAC ACG T GCA AGA R ATA CTT GGA G TGT CAG $T^{T}C$ TGT C AGG R TCT ATT I TTTATG M CGA ACA GAC D GTT V CTT GAA E TCTAGA R TTG AAT L N CAA Q CGT R ggtctcgagacttcagcaatattgacacatcag G TTC F AAC GTT AG R AGG R CCC P CTTAGT S CCA AGA TTA ATTД aaagtcattaattaaccttag AT ACG T CCC TTT CAT CTA TAT \mathtt{TAT} TAC AAA K AAG K GAT GAT D GGA G ATC ATT AAG K GAA ATA E I AAA GAT ATG M TCTCTTΩ CCT AGG R CCC GATAAA AGC GCT 4210 889

F/G. 15 (CONTINUED)

 Atty. Docket No.: 015389-002640US Applicant: Thomas R. Cech et al.

DMÉRASE CATALYTIC SUBUNIT: DIAGNOSTIC A

THERAPEUTIC METHODS

Sheet 19 of 34

19/34

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ttatccttatacttttaagaaagattgacagtggttgctgactactgcccacatgcccattaaacgggagtggttaaaca ttaaaagtaatacatgaggctaatctcctttcatttagaataaggaaagtggttttctataatgaataatgcccgcacta

> 4986 4826 4906

5066

atgcaaaaagacgaagattatcttctaaacaagggggattaagcatatccgaaggaaaagagagattaatatacccagtgtt gttgaagaaagcaaggataatttggaacaagcttctgcagatgacaggctaaattttggtgaccgaattttggtaaaagc toctgatttaaaaggaggaatcttccaccgatgaggaaatggatagcttatcagctgctgaggagaagcctaatttttgc cccaggttatccatggtggccggccttgctactgagacgaaaagaaactaaggatagtttgaatactaatagctcattta atgictiatataaaggittigittitticcigacticaatittigcaiggigaaaagaaatagigtiaagccattaitggat tecgaaatagecaaatttettggtteeteaaageggaagtetaaagaaettattgaagettatgaggetteaaaaaetee aaaaaagaaaatatcattgggagacatctcttgatgaatcagatgcggagagtatctccagcggatccttgatgtcaata acttctatttctgaaatgtatggtcctactgtcgcttcgacttctcgtagctctacgcagttaagtgaccaaaggtacc (CONTINUED) FIG. 15

 Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
Title: HUMAN DMERASE CATALYTIC SUBUNIT: DIAGNOSTIC ATTHERAPEUTIC METHODS
Sheet 20 of 34

20/34

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	gcgcgctccc					
	gccgctggcc					
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241	acggccgccc	cccgccgccc	cctccttccg	ccaggtgtcc	tgcctgaagg	agctggtggc
301	ccgagtgctg	cagaggctgt	acaaacacaa	cqcqaaqaac	gtgctggcct	teggettege
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541	ggctcccagc	tgcgcctacc	aggtgtgcgg	gccgccgctg	taccageteg	gcgctgccac
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781	tgcccctgag	ccggagcgga	cgcccgttgg	gcaggggtcc	tgggcccacc	cgggcaggac
841	gcgtggaccg	agtgaccgtg	gtttctgtgt	ggtgtcacct	gccagacccg	ccgaagaagc
	cacctctttg					
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1001	gcaccacgcg	ggccccccac	ccacaccgcg	gccaccacgc	cccgggaca	cgccccgccc
	cccggtgtac					
1081	gccctccttc	ctactcagct	ctctgaggcc	cagcctgact	ggcgctcgga	ggctcgtgga
1141	gaccatcttt	ctgggttcca	ggccctggat	gccagggact	ccccgcaggt	tgccccgcct
	gccccagcgc					
	gtgcccctac					
1321	agccggtgtc	tatacccada	agaagcccca	gggctctgtg	geggeeeeeg	aggaggagga
1381	cacagacccc	cgtcgcctgg	tgcagctgct	ccgccagcac	agcagcccct	ggcaggtgta
1441	cggcttcgtg	caaacctacc	tacaccaact	ggtgccccca	ggcctctggg	gctccaggca
1501	caacgaacgc	cocttectea	ggaacaccaa	gaagttcatc	tecetagaga	agcatgccaa
1561	gctctcgctg	caccacctca	cataassast	gaageteaa	gactgcgctt	agetacacaa
	gagcccaggg					
	caagttcctg					
1741	tgtcacggag	accacgtttc	aaaagaacag	gctctttttc	taccggaaga	gtgtctggag
	caagttgcaa					
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	cttcatcccc					
1981	cagaacgttc	cgcagagaaa	agagggccga	gcgtctcacc	tcgagggtga	aggcactgtt
2041	cagcgtgctc	aactacgagc	gggcgcggcg	ccccggcctc	ctgggcgcct	ctgtgctggg
	cctggacgat					
	gccgcctgag					
	ggacaggctc					
2281	tcggtatgcc	gtggtccaga	aggccgccca	tgggcacgtc	cgcaaggcct	tcaagagcca
2341	cgtctctacc	ttgacagacc	tccagccgta	catgcgacag	ttcgtggctc	acctgcagga
	gaccagcccg					
2/61	cagtggcctc	ttccacatct	tectaceett	catotoccac	caccottoc	acatcadad
2501	caguagecte	atasatasa	ccccacgccc	catgegetae	cacgeege	gcaccagggg
Z3Z1	caagtcctac	gcccagcgcc	aggggateee	geagggeree	accouctoca	egetgetetg
	cagcctgtgc					
2641	gctcctgcgt	ttggtggatg	atttcttgtt	ggtgacacct	cacctcaccc	acgcgaaaac
2701	cttcctcagg	accetagtee	gaggtgtccc	tgagtatggc	tacataataa	acttgcggaa
2761	gacagtggtg	aacttccctc	tagaagagg	agecetagat	agracagett	ttattcagat
2021	gacageggeg	contatta	cagaagacga	ggccccgggc	ggcacggccc	cogecoagae
2021	gccggcccac	ggeeratiee	ceragingeng	cetgetgetg	gatacccyga	ccctggaggt
	gcagagcgac					
2941	cggcttcaag	gctgggagga	acatgcgtcg	caaactcttt	ggggtcttgc	ggctgaagtg
	tcacagcctg					
3061	caagatcctc	ctactacaaa	catacadatt	tracgratut	atactacaac	tcccatttca
	tcagcaagtt					
3181	ctgctactcc	atcctgaaag	ccaagaacgc	agggatgtcg	ctgggggcca	agggcgccgc
3241	cggccctctg	ccctccgagg	ccgtgcagtg	gctgtgccac	caagcattcc	tgctcaagct
3301	gactcgacac	catatcacct	acqtqccact	cctagaatca	ctcaggacag	cccagacaca
3361	gctgagtcgg	aanctccccc	udacdacdct	gactgccctg	daddccdcad	ccaacccccc
3/01	act canter	anguicecyy	ggacgacgcc	atantaara	adagaaaaaa	accade course
	actgccctca					
3481	gagcagacac	cagcagccct	gtcacgccgg	gctctacgtc	ccagggaggg	aggggcggcc
3541	cacacccagg	cccgcaccgc	tgggagtctg	aggcctgagt	gagtgtttgg	ccgaggcctg
3601	catgtccggc	tgaaggctga	atatecaact	gaggcctgag	cgagtgtcca	gccaagggct
3661	gagtgtccag	cacacctacc	atattasatt	CCCCSCSCCC	taacaataaa	ctccacccca
2771	avacac	thank	gue coude	thaaaataa	caacacccaa	tartacataa
7/27	gggccagctt	LLCCTCacca	yyaycccggc	LLCCACCCCC	cacacaggaa	Lagicocatoc
3/81	ccagattcgc	cattgttcac	ccctcgccct	gccctccttt	gccttccacc	cccaccatcc
3841	aggtggagac	cctgagaagg	accctgggag	ctctgggaat	ttggagtgac	caaaggtgtg
3901	ccctgtacac	aggcgaggac	cctgcacctg	gatgggggtc	cctataaata	aaattaaaaa
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MPRAPRCRAVRSLLRSHYREVLPLATFVRRLGPQGWRLVQRGDP AAFRALVAQCLVCVPWDARPPPAAPSFRQVSCLKELVARVLQRL CERGAKNVLAFGFALLDGARGGPPEAFTTSVRSYLPNTVTDALR GSGAWGLLLRRVGDDVLVHLLARCALFVLVAPSCAYQVCGPPLY OLGAATQARPPPHASGPRRRLGCERAWNHSVREAGVPLGLPAPG ARRRGGSASRSLPLPKRPRRGAAPEPERTPVGQGSWAHPGRTRG PSDRGFCVVSPARPAEEATSLEGALSGTRHSHPSVGRQHHAGPP STSRPPRPWDTPCPPVYAETKHFLYSSGDKEQLRPSFLLSSLRP SLTGARRLVETIFLGSRPWMPGTPRRLPRLPQRYWQMRPLFLEL LGNHAQCPYGVLLKTHCPLRAAVTPAAGVCAREKPQGSVAAPEE EDTDPRRLVQLLRQHSSPWQVYGFVRACLRRLVPPGLWGSRHNE RRFLRNTKKFISLGKHAKLSLQELTWKMSVRDCAWLRRSPGVGC VPAAEHRLREEILAKFLHWLMSVYVVELLRSFFYVTETTFQKNR LFFYRKSVWSKLQSIGIRQHLKRVQLRELSEAEVRQHREARPAL LTSRLRFIPKPDGLRPIVNMDYVVGARTFRREKRAERLTSRVKA LFSVLNYERARRPGLLGASVLGLDDIHRAWRTFVLRVRAQDPPP ELYFVKVDVTGAYDTIPQDRLTEVIASIIKPQNTYCVRRYAVVQ KAAHGHVRKAFKSHVSTLTDLQPYMRQFVAHLQETSPLRDAVVI EQSSSLNEASSGLFDVFLRFMCHHAVRIRGKSYVQCQGIPQGSI LSTLLCSLCYGDMENKLFAGIRRDGLLLRLVDDFLLVTPHLTHA KTFLRTLVRGVPEYGCVVNLRKTVVNFPVEDEALGGTAFVQMPA HGLFPWCGLLLDTRTLEVQSDYSSYARTSIRASLTFNRGFKAGR NMRRKLFGVLRLKCHSLFLDLQVNSLQTVCTNIYKILLLQAYRF HACVLQLPFHQQVWKNPTFFLRVISDTASLCYSILKAKNAGMSL GAKGAAGPLPSEAVQWLCHQAFLLKLTRHRVTYVPLLGSLRTAQ TQLSRKLPGTTLTALEAAANPALPSDFKTILD

FIG. 17

TTATGTCACGGAGACCACGTTTCAAAAGAACAGGCTCTTTTTCTACCGGAAGAGTGTCTG GAGCAAGTTGCAAAGCATTGGAATCAGACAGCACTTGAAGAGGGTGCAGCTGCGGGAGCT CCGCTTCATCCCCAAGCCTGACGGCCTGCGGCCGATTGTGAACATGGACTACGTCGTGGG AGCCAGAACGTTCCGCAGAGAAAAGAGGGCCGAGCGTCTCACCTCGAGGGTGAAGGCACT GTTCAGCGTGCTCAACTACGAGCGGGCGCGCGCCCCGGCCTCCTGGGCGCCTCTGTGCT GGGCCTGGACGATATCCACAGGGCCTGGCGCACCTTCGTGCTGCGTGTGCGGGCCCAGGA CCCGCCGCCTGAGCTGTACTTTGTCAAGGTGGATGTGACGGGCGCGTACGACACCATCCC CCAGGACAGGCTCACGGAGGTCATCGCCAGCATCATCAAACCCCAGAACACGTACTGCGT GCGTCGGTATGCCGTGGTCCAGAAGGCCGCCCATGGGCACGTCCGCAAGGCCTTCAAGAG CCACGTCCTACGTCCAGTGCCAGGGGATCCCGCAGGGCTCCATCCTCTCCACGCTGCTCT GCAGCCTGTGCTACGGCGACATGGAGAACAAGCTGTTTGCGGGGGATTCGGCGGGACGGGC TGCTCCTGCGTTTGGTGGATGATTTCTTGTTGGTGACACCTCACCTCACCCACGCGAAAA CCTTCCTCAGGACCCTGGTCCGAGGTGTCCCTGAGTATGGCTGCGTGAACTTGCGGA AGACAGTGGTGAACTTCCCTGTAGAAGACGAGGCCCTGGGTGGCACGGCTTTTGTTCAGA TGCCGGCCCACGGCCTATTCCCCTGGTGCGGCCTGCTGCTGGATACCCGGACCCTGGAGG TGCAGAGCGACTACTCCAGCTATGCCCGGACCTCCATCAGAGCCAGTCTCACCTTCAACC GCGGCTTCAAGGCTGGGAGGAACATGCGTCGCAAACTCTTTGGGGTCTTGCGGCTGAAGT GTCACAGCCTGTTTCTGGATTTGCAGGTGAACAGCCTCCAGACGGTGTGCACCAACATCT ACAAGATCCTCCTGCTGCAGGCGTACAGGTTTCACGCATGTGTGCTGCAGCTCCCATTTC ATCAGCAAGTTTGGAAGAACCCCACATTTTTCCTGCGCGTCATCTCTGACACGGCCTCCC TCTGCTACTCCATCCTGAAAGCCAAGAACGCAGGGATGTCGCTGGGGGCCAAGGGCGCCG CCGGCC7TCTGCCCTCCGAGGCCGTGCAGTGGCTGTGCCACCAAGCATTCCTGCTCAAGC TGACTCGACACCGTGTCACCTACGTGCCACTCCTGGGGTCACTCAGGACAGCCCAGACGC AGCTGAGTCGGAAGCTCCCGGGGACGACGCTGACTGCCCTGGAGGCCGCAGCCAACCCGG CACTGCCCTCAGACTTCAAGACCATCCTGGACTGATGGCCACCCGCCCACAGCCAGGCCG GCATGTCCGGCTGAAGGCTGAGTGTCCGGCTGAGGCCTGAGCGAGTGTCCAGCCAAGGGC TGAGTGTCCAGCACCTGCCGTCTTCACTTCCCCACAGGCTGGCGCTCGGCTCCACCCC AGGGCCAGCTTTTCCTCACCAGGAGCCCGGCTTCCACTCCCCACATAGGAATAGTCCATC CCCAGATTCGCCATTGTTCACCCCTCGCCCTGCCCTCCTTTGCCTTCCACCCCCACCATC CAGGTGGAGACCCTGAGAAGGACCCTGGGAGCTCTGGGAATTTGGAGTGACCAAAGGTGT GCCTGTACACAGGCGAGGACCCTGCACCTGGATGGGGGTCCCTGTGGGTCAAATTGGGG AAAAAAAAAAAAA

Atty. Docket No.: 015389-002640US Applicant: Thomas R. Cech et al. Title: HUMAN LOMERASE CATALYTIC SUBUNIT: DIAGNOSTIC THERAPEUTIC METHODS Sheet 22 of 34

MetSerValTyrValValGluLeuLeuArgSerPhePhe TyrValThrGluThrThrPheGlnLysAsnArgLeuPhe PheTyrArgLysSerValTrpSerLysLeuGlnSerIle GlyIleArgGlnHisLeuLysArgValGlnLeuArgGlu LeuSerGluAlaGluValArgGlnHisArgGluAlaArg ProAlaLeuLeuThrSerArgLeuArgPheIleProLys ProAspGlyLeuArgProIleValAsnMetAspTyrVal ValGlyAlaArgThrPheArgArgGluLysArgAlaGlu ArgLeuThrSerArgValLysAlaLeuPheSerValLeu AsnTyrGluArgAlaArgArgProGlyLeuLeuGlyAla SerValLeuGlyLeuAspAspIleHisArgAlaTrpArg ThrPheValLeuArgValArgAlaGlnAspProProPro ${\tt GluLeuTyrPheValLysValAspValThrGlyAlaTyr}$ AspThrIleProGlnAspArgLeuThrGluValIleAla SerIleIleLysProGlnAsnThrTyrCysValArgArg TyrAlaValValGlnLysAlaAlaHisGlyHisValArg LysAlaPheLysSerHisValLeuArgProValProGly AspProAlaGlyLeuHisProLeuHisAlaAlaLeuGln ProValLeuArgArgHisGlyGluGlnAlaValCysGly AspSerAlaGlyArgAlaAlaProAlaPheGlyGly

GCA	GCGC'	rgcgʻ	rccro	GCTG	CGCA	CGTG	GGAA(geee:	rggc	cccc	GCCA	cccc	CGCG	1 met ATG
								10	•.					
								val				leu CTG		
												arg CGG		
												pro CCG		
												pro		
												val GTG		
												CYS TGC		
												leu CTG		
ala GCC	arg CGC	gly GGG	110 gly GGC	pro CCC	pro CCC	glu GAG	ala GCC	phe TTC	thr ACC	thr ACC	ser AGC	val GTG	120 arg CGC	ser AGC

Atty. Docket No.: 015389-002640US Applicant: Thomas R. Cech et al. Title: HUMA LOMERASE CATALYTIC SUBUNIT: DIAGNOSTIC THERAPEUTIC METHODS Sheet 23 of 34

		pro CCC												
		leu CTG												
leu CTG	leu CTG	ala GCA	arg CGC	cys TGC	ala GCG	leu CTC	phe TTT	160 val GTG	leu CTG	val GTG	ala GCT	pro CCC	ser AGC	cys TGC
		gln CAG												
		ala GCC												
		cys TGC												
val GTC	pro CCC	leu CTG	gly GGC	leu CTG	pro CCA	ala GCC	pro CCG	220 gly GGT	ala GCG	arg AGG	arg AGG	arg CGC	gly GGG	gly GGC
		ser AGC												
		pro CCT												
		pro CCG												
val GTG	val GTG	ser TCA	pro CCT	ala GCC	arg AGA	pro CCC	ala GCC	280 glu GAA	glu GAA	ala GCC	thr ACC	ser TCT	leu TTG	glu GAG
gly GGT	ala GCG	leu CTC	290 ser TCT	gly	thr ACG	arg CGC	his CAC	ser TCC	his CAC	pro CCA	ser TCC	val GTG	300 gly GGC	arg CGC
		his CAC												pro CCC
trp TGG	asp GAC	thr ACG	320 pro CCT	cys TGT	pro CCC	pro CCG	val GTG	tyr TAC	ala GCC	glu GAG	thr ACC	lys AAG	330 his CAC	phe TTC

FIG. 20 (CONTINUED)

Atty. Docket No.: 015389-002640US Applicant: Thomas R. Cech et al. Title: HUMAN THERASE CATALYTIC SUBUNIT: DIAGNOSTIC AND THERAPEUTIC METHODS Sheet 24 of 34

	leu CTC	tyr TAC	ser TCC	ser TCA	gly GGC	asp GAC	lys AAG	glu	gln CAG	leu CTG	arg CGG	pro CCC	ser TCC	phe TTC	leu CTA
-	leu CTC	ser AGC	ser TCT	350 leu CTG	arg AGG	pro CCC	ser AGC	leu CTG	thr ACT	gly GGC	ala GCT	arg CGG	arg AGG	360 leu CTC	val GTG
(glu GAG	thr ACC	ile ATC	phe TTT	leu CTG	gly GGT	ser TCC	arg AGG	370 pro CCC	trp TGG	met ATG	pro CCA	gly GGG	thr ACT	pro CCC
ä	arg CGC	arg AGG	leu TTG	380 pro	arg CGC	leu CTG	pro CCC	gln CAG	arg CGC	tyr TAC	trp TGG	gln CAA	met ATG	390 arg CGG	pro CCC
	leu CTG	phe TTT	leu CTG	glu GAG	leu CTG	leu CTT	GGG Gly	asn AAC	400 his CAC	ala GCG	gln CAG	cys TGC	pro CCC	tyr TAC	gly GGG
	val GTG	leu CTC	leu CTC	410 lys AAG	thr ACG	his CAC	cys TGC	pro CCG	leu CTG	arg CGA	ala GCT	ala GCG	val GTC	420 thr ACC	pro CCA
	ala GCA	ala GCC	gly GGT	val GTC	cys TGT	ala GCC	arg CGG	glu GAG	430 lys AAG	pro	gln CAG	gly GGC	ser TCT	val GTG	ala GCG
,	ala GCC	pro CCC	glu GAG	440 glu GAG	glu GAG	asp GAC	thr ACA	asp GAC	pro CCC	arg CGT	arg CGC	leu CTG	val GTG	450 gln CAG	leu CTG
	leu CTC	arg CGC	gln CAG	his CAC	ser AGC	ser AGC	pro CCC	trp TGG	460 gln CAG	val GTG	tyr TAC	gly GGC	phe TTC	val GTG	arg CGG
	ala GCC	cys TGC	leu CTG	470 arg CGC	arg CGG	leu CTG	val GTG	pro CCC	pro CCA	gly GGC	leu CTC	trp TGG	gly GGC	480 ser TCC	arg AGG
	his CAC	asn AAC	glu GAA	arg CGC	arg CGC	phe TTC	leu CTC	arg AGG	490 asn AAC	thr ACC	lys AAG	lys AAG	phe TTC	ile ATC	ser TCC
	leu CTG	gly GGG	lys AAG	500 his CAT	ala GCC	lys AAG	leu CTC	ser TCG	leu CTG	gln CAG	glu GAG	leu CTG	thr ACG	510 trp TGG	lys AAG
	met ATG	ser AGC	val GTG	arg CGG	asp GAC	cys TGC	ala GCT	trp TGG	520 leu CTG	arg CGC	arg AGG	ser AGC	pro CCA	gly GGG	val GTT
	gly GGC	cys TGT	val GTT	530 pro CCG	ala GCC	ala GCA	glu GAG	his CAC	arg CGT	leu CTG	arg CGT	glu GAG	glu GAG	540 ile ATC	leu CTG

FIG. 20 (CONTINUED)

Atty. Docket No.: 015389-002640US Applicant: Thomas R. Cech et al. LOMERASE CATALYTIC SUBUNIT: DIAGNOSTIC THERAPEUTIC METHODS Sheet 25 of 34

							5	550						
		phe TTC												
		ser TCT												
arg AGG	leu CTC	phe TTT	phe TTC 590	tyr TAC	arg CGG	lys AAG	ser AGT	580 val GTC	trp TGG	ser AGC	lys AAG	leu TTG	gln CAA 600	ser AGC
ile ATT	gly GGA	ile ATC	arg AGA	gln CAG	his CAC	leu TTG	lys AAG	arg AGG	val GTG	gln CAG	leu CTG	arg CGG	glu GAG	leu CTG
ser TCG	glu GAA	ala GCA	glu GAG	val GTC	arg AGG	gln CAG	his CAT	610 arg CGG	glu GAA	ala GCC	arg AGG	pro CCC	ala GCC	leu CTG
leu CTG	thr ACG	ser TCC	620 arg AGA	leu CTC	arg CGC	phe TTC	ile ATC	pro CCC	lys AAG	pro CCT	asp GAC	gly GGG	630 leu CTG	arg CGG
pro CCG	ile ATT	val GTG	asn AAC	met ATG	asp GAC	tyr TAC	val GTC	640 val GTG	gly GGA	ala GCC	arg AGA	thr ACG	phe TTC	arg
arg AGA	glu GAA	lys AAG	650 arg AGG	ala GCC	glu GAG	arg CGT	leu CTC	thr ACC	ser TCG	arg AGG	val GTG	lys AAG	660 ala GCA	leu CTG
phe TTC	ser AGC	val GTG	leu CTC	asn AAC	tyr TAC	glu GAG	arg CGG	670 ala GCG	arg CGG	arg CGC	pro	gly GGC	leu CTC	leu CTG
gly GGC	ala GCC	ser TCT	680 val GTG	leu CTG	gly GGC	leu CTG	asp GAC	asp GAT	ile ATC	his CAC	arg AGG	ala GCC	690 trp TGG	arg CGC
thr ACC	phe TTC	val GTG	leu CTG	arg CGT	val GTG	arg CGG	ala GCC	700 gln CAG	asp GAC	pro CCG	pro CCG	pro CCT	glu GAG	leu CTG
tyr TAC	phe TTT	val GTC	710 lys AAG	val GTG	asp GAT	val GTG	thr ACG	gly GGC	ala GCG	tyr TAC	asp GAC	thr ACC	720 ile ATC	pro CCC
gln CAG	asp GAC	arg AGG	leu CTC	thr ACG	glu GAG	val GTC	ile ATC	730 ala GCC	ser AGC	ile ATC	ile ATC	lys AAA	pro CCC	gln CAG
asn AAC	thr ACG	tyr TAC	740 cys TGC	val GTG	arg CGT	arg CGG	tyr TAT	ala GCC	val GTG	val GTC	gln CAG	lys AAG	750 ala GCC	ala GCC

FIG. 20 (CONTINUED)

760 his gly his val arg lys ala phe lys ser his val leu arg pro CAT GGG CAC GTC CGC AAG GCC TTC AAG AGC CAC GTC CTA CGT CCA 780 770 val pro gly asp pro ala gly leu his pro leu his ala ala leu GTG CCA GGG GAT CCC GCA GGG CTC CAT CCT CTC CAC GCT GCT CTG 790 gln pro val leu arg arg his gly glu gln ala val cys gly asp CAG CCT GTG CTA CGG CGA CAT GGA GAA CAA GCT GTT TGC GGG GAT 807 800 ser ala gly arg ala ala pro ala phe gly gly OP TCG-GCG-GGA-CGG-GCT-GCT-GCG-TTT-GGT-GGA-TGA-TTTCTTGTTGGT GACACCTCACCTCACCCACGCGAAAACCTTCCTCAGGACCCTGGTCCGAGGTGTCCCTGA GTATGGCTGCGTGAACTTGCGGAAGACAGTGGTGAACTTCCCTGTAGAAGACGAGGC CCTGGGTGGCACGGCTTTTGTTCAGATGCCGGCCCACGGCCTATTCCCCTGGTGCGGCCT GCTGCTGGATACCCGGACCCTGGAGGTGCAGAGCGACTACTCCAGCTATGCCCGGACCTC CATCAGAGCCAGTCTCACCTTCAACCGCGGCTTCAAGGCTGGGAGGAACATGCGTCGCAA ACTCTTTGGGGTCTTGCGGCTGAAGTGTCACAGCCTGTTTCTGGATTTGCAGGTGAACAG CCTCCAGACGTGTGCACCAACATCTACAAGATCCTCCTGCTGCAGGCGTACAGGTTTCA CGCATGTGTGCTGCAGCTCCCATTTCATCAGCAAGTTTGGAAGAACCCCACATTTTTCCT GCGCGTCATCTCTGACACGGCCTCCCTCTGCTACTCCATCCTGAAAGCCAAGAACGCAGG GATGTCGCTGGGGGCCAAGGGCGCCGCCGGCCTCTGCCCTCCGAGGCCGTGCAGTGGCT GTGCCACCAAGCATTCCTGCTCAAGCTGACTCGACACCGTGTCACCTACGTGCCACTCCT GGGGTCACTCAGGACAGCCCAGACGCAGCTGAGTCGGAAGCTCCCGGGGACGACGCTGAC TGCCCTGGAGGCCGCAGCCAACCCGGCACTGCCCTCAGACTTCAAGACCATCCTGGACTG CTACGTCCCAGGGAGGGGGGGCCCCACACCCAGGCCCGCACCGCTGGGAGTCTGAGG CCTGAGTGAGTGTTTGGCCGAGGCCTGCATGTCCGGCTGAAGGCTGAGTGTCCGGCTGAG GCCTGAGCGAGTGTCCAGCCAAGGGCTGAGTGTCCAGCACACCTGCCGTCTTCACTTCCC CACAGGCTGGCGCTCCACCCCAGGGCCAGCTTTTCCTCACCAGGAGCCCGGCTTC CACTCCCCACATAGGAATAGTCCATCCCCAGATTCGCCATTGTTCACCCCTCGCCCTGCC CTCCTTTGCCTTCCACCCCCACCATCCAGGTGGAGACCCTGAGAAGGACCCTGGGAGCTC TGGGAATTTGGAGTGACCAAAGGTGTGCCCTGTACACAGGCGAGGACCCTGCACCTGGAT GGGGGTCCCTGTGGGTCAAATTGGGGGGGGGGTGCTGTGGGAGTAAAATACTGAATATATG

> FIG. 20 (CONTINUED)

3601	1 ATCGATTGGGCCCGAGATCTCGCGCGCGCGAGGCCTGCCATGGGACCCAC TAGCTAACCCGGGCTCTAGAGCGCGCGCTCCGGACGTACCCTGGGTG.	
	3615 3636 BGL2 NCO1	
3661	TGGGANGCTGCAGGCTTCAGGTCCCAGTGGGGTTGCCATCTGCCAGTAGACCTNCGACGTCCGAAGTCCAGGGTCACCCCAACGGTAGACGGTCATCATC	
3721	21 AGAATCAGGGCGCGAGTGTGGACACTGTCCTGAATCTCAATGTCTCAG TCTTAGTCCCGCGCTCACACCTGTGACAGGACTTAGAGTTACAGAGTC.	
3781	31 CATGTAGAAATTAAAGTCCATCCTCCTACTCTACTGGGATTGAGCCCGGGTACATCTTTAATTTCAGGTAGGGAGGATGAGATGACCCTAACTCGGG	
3841	11 CCCCCAGGGGCAGAGGAGTTCCTCTCACTCCTGTGGAGGAAGGA	
3901	1 TTTCACTGCTGGTACTGAATCCACTGTTTCATTTGTTGGTTTGTTGT AAAGTGACGACCATGACTTAGGTGACAAAGTAAACAACCAAACAAA	TTTGTTTTGAGA AAACAAAACTCT
3961	**************************************	TCTTGGCTTACT
	ALU ************************************	*****
4021	21 GCAGCCTCTGCCTCCCAGGTTCAAGTGATTCTCCTGCTTCCGCCTCCC CGTCGGAGACGGAGGGTCCAAGTTCACTAAGAGGACGAAGGCGGAGGG	ATTTGGCTGGGA
	******	*****
4081	31 TTACAGGCACCCGCCACCATGCCCAGCTAATTTTTTGTATTTTAGTA AATGTCCGTGGGCGGTGGTACGGGTCGATTAAAAAACATAAAAATCAT	
		A
4141	11 GGGGTGGGGTTCACATGTTGGCCAAGCTGGTCTCGAACTTCTGAACTC CCCCACCCCA	
	LU	
4201	TGCCTCTGCCTCTAAAATTGCTGGGATTACAGGTGTNANCCACCATG ACGGAGACGGAGATTTAACGACCCTAATGTCCACANTNGGTGGTAC	CCCAACTCAAAA
4261	TTTACTCTGTTTANAAACATCTGGGTCTAAGGTAGGAANCTCACCCCA AAATGAGACAAATNTTTGTAGACCCAGATTCCATCCTTNGAGTGGGGT	

432,1	GGTGTTTTTAAGCCAATNANAAAATTTTTTTNATGTTGTTTNNNNNNNNNN
4381	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4441	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4501	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4561	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4621	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4681	ИПИМИНИИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИ
4741	ИИМИИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4801	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4861	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4921	ИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМИМ
4981	NNINNNINNNINNNINNNINNNINNNINNNINNNINNN
5041	NGCCANGRAGGGGCCAGGTTCCAANTTCCCAACCKTTTTWGGARGGACNGCCCCCAGGG NCGGTNCYTCCCCCGGTCCAAGGTTNAAGGGTTGGMAAAAWCCTYCCTGNCGGGGGTCCC
5101	GGGGATRAACAGANTNGGGGGKGGTWGGGTTNAKGGTGGGAACNCCTTNGCGCCTGGAG CCCCTAYTTGTCTNANCCCCCMCCAWCCCAANTMCCACCCTTGNGGAANCGSCGGACCTC
5161	AACGTGCAAAGAGGAAATGAAGGGCCTGKGTCAAGGAGCCCAAGTNGGCGGGGRAGTTTG TTGCACGTTTCTCCTTTACTTCCCGGACMCAGTTCCTCGGGTTCANCCGCCCCYTCAAAC
5221	CAGGGAGGCACTCCGGGGAGGTCCSGCGTGCCCGTCCAAGGGAGCAATGCGTCCTTCGGG GTCCCTCCGTGAGGCCCCTCCAGGSCGCACGGGCAGGTTCCCTCGTTACGCAGGAAGCCC
5281	TTCGTCCCCAWGCCGCGTCTACGCGCCTYCCGTCCTCCCCTTCACGTTCCGGCATTCGTG AAGCAGGGGTWCGGCGCAGATGCGCGGARGGCAGGAGGGGAAGTGCAAGGCCGTAAGCAC
5341	GTGCCCGGAGCCCGACGCCCCGCGTCCGGACCTGGAGCCAGGCCCTGGGTCTCCGGATCAGCACGGGCCTCGGGCCTCGGGCCTAGTCCACGGGCCTCGGGCCCAGAGGCCTAGTC
5401	GCCAGCGGCCAAAGGGTCGCCGCACGCACCTGTTCCCAGGGCCTCCACATCATGGCCCCTCGGTCGCCGGTTTCCCAGCGCGCGTGCGT

FIG. 21 (CONTINUED)

5461 CCCTCGGGTTACCCCACAGCCTAGGCCGGATTCGACCTCTCTCCGCTGGGGCCCTCGCCT GGGAGCCCAATGGGGTGTCGGATCCGGCCTAAGCTGGAGAGAGGCGACCCCGGGAGCGGA Sp1 CCGCAGGGACGTGGGACCCTCGCGCTCGCCGCGCGCCCCCTTCGCGCCGGGTATGGG 5581 CCGGGTCCGCCGGAAGCAGCTGCGCTGTCGGGGCCAGGCCGGGCTCCCAGTGGATTCGC GGCCAGGCGGCCTTCGTCGACGCGACAGCCCCGGTCCGGCCCGAGGGTCACCTAAGCG Topo_II_cleavage_site 5641 GGGCACAGACGCCCAGGACCGCGCTTCCCACGTGGCGAAGGACTGGGGACCCGGGCACC CCCGTGTCTGCGGGTCCTGGCGCGAAGGGTGCACCGCCTTCCTGACCCCTGGGCCCGTGG E2F ***** 5701 CGTCCTGCCCCTTCACCTTCCAGCTCCGCTTCTTCCGCGCGGACCCGGCCCCGTCCCGAA GCAGGACGGGGAAGTGGAAGGTCGAGGCGAAGAAGGCGCGCCTGGGCCGGGGCAGGGCTT E Sp1 ======= h 2F NFkB **** 5821 CGCGGCCCCGCCTCTCCTTCGCGGCGCGAGTTTCAGGCAGCGCTGCGTCCTGCTGCGCA GCGCCGGGGCGGAGAGGAAGCGCCGCGCTCAAAGTCCGTCGCGACGCAGGACGACGCGT 5860 5875 ECO47III FSP1 ******* 5881 CGTGGGAAGCCCTGGCCCGGCCACCCCGCGATGCCGCGCTCCCCGCTGCCGAGCCG GCACCCTTCGGGACCGGGGCCGGTGGGGGGCGCTACGGCGCGCGAGGGGCGACGGCTCGGC 5941 TGCGCTCCCTGCTGCGCAGCCACTACCGCGAGGTGCTGCCGCTGGCCACGTTCGTGCGGC ACGCGAGGGACGACGCTCGTGATGCCGCTCCACGACGGCGACCGGTGCAAGCACGCCG 5953

NFkB

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FIG. 21 (CONTINUED)

*********** 6121 CCTTCCGCCAGGTGGGCCTCCCCGGGGTCGGCGTCCGGCTGGGGTTGAGGGCGGCCGGGG GGAAGGCGGTCCACCCGGAGGGGCCCCAGCCGCAGCCCCAACTCCCGCCGGCCCC Topo_II_cleavage_s NFkB ========= Intron1 ************** 6181 GGAACCAGCGACATGCGGAGAGCAGCGCAGGCGACTCAGGGCGCTTCCCCCGCAGGTGTC CCTTGGTCGCTGTACGCCTCTCGTCGCGTCCGCTGAGTCCCGCGAAGGGGGCGTCCACAG ite 6301 CGTGCTGGCCTTCGCCTTCGCGCTGCTGGACGGGGCCCCCCGGGGGCCCCCCGAGGCCTT GCACGACCGGAAGCCGAAGCGCGACGACCTGCCCCGGGCGCCCCCGGGGGGGCTCCGGAA 6361 CACCACCAGCGTGCGCAGCTACCTGCCCAACACGGTGACCGACGCACTGCGGGGGAGCGG GTGGTGGTCGCACGCGTCGATGGACGGGTTGTGCCACTGGCTGCGTGACGCCCCCTCGCC 6372 FSP1 6421 GGCGTGGGGGCTGCTGCGCCGCGTGGGCGACGACGTGCTGGTTCACCTGCTGGCACG CCGCACCCCGACGACGACGCGCGCCCCCGCTGCACGACCAAGTGGACGACCGTGC 6481 CTGCGCGCTCTTTGTGCTGGTGGCTCCCAGCTGCGCCTACCAGGTGTGCGGGCCGCCGCT GACGCGCGAGAAACACGACCACCGAGGGTCGACGCGGATGGTCCACACGCCCGGCGGCGA 6541 GTACCAGCTCGGCGCTGCCACTCAGGCCCGGCCCCGCCACACGCTAGTGGACCCCGAAG 6601 GCGTCTGGGATGCGAACGGGCCTGGAACCATAGCGTCAGGGAGGCCGGGGTCCCCCTGGG CGCAGACCCTACGCTTGCCCGGACCTTGGTATCGCAGTCCCTCCGGCCCCAGGGGGACCC 6661 CCTGCCAGCCCGGGTGCGAGGAGGCGCGGGGGCAGTGCCAGCCGAAGTCTGCCGTTGCC GGACGGTCGGGGCCCACGCTCCTCCGCGCCCCCGTCACGGTCGGCTTCAGACGGCAACGG 6721 CAAGAGGCCCAGGCGTGCCCCTGAGCCGGAGCGGACGCCCGTTGGGCAGGGGTC GTTCTCCGGGTCCGCACCGCGACGGGGACTCGCCTGCCGGGCAACCCGTCCCCAG GACCCGGGTGGGCCCGTCCTGCGCACCTGGCTCACTGGCACCAAAGACACACCACAGTGG 6841 TGCCAGACCCGCGAAGAAGCCACCTCTTTGGAGGGTGCGCTCTCTGGCACGCGCCACTC ACGGTCTGGGCGGCTTCTTCGGTGGAGAAACCTCCCACGCGAGAGACCCGTGCGCGGTGAG

FIG. 21 (CONTINUED)

GGTGGGTAGCCCCGGCGGTCGTGCTGCGCCCGGGGGTAGGTGTAGCGCCGGTGGTGC

6901 CCACCCATCCGTGGGCCGCCAGCACCACGCGGGCCCCCCATCCACATCGCGGCCACCACG

- 6961 TCCCTGGGACACGCCTTGTCCCCCGGTGTACGCCGAGACCAAGCACTTCCTCTACTCCTC AGGGACCCTGTGCGGAACAGGGGGCCACATGCGGCTCTGGTTCGTGAAGGAGATGAGGAG
- 7021 AGGCGACAAGGAGCAGCTGCGGCCCTCCTTCCTACTCAGCTCTCTGAGGCCCAGCCTGAC
 TCCGCTGTTCCTCGTCGACGCCGGGAGGAAGGATGAGTCGAGAGACTCCGGGTCGGACTG
- 7081 TGGCGCTCGGAGGCTCGTGGAGACCATCTTTCTGGGTTCCAGGCCCTGGATGCCAGGGAC
 ACCGCGAGCCTCCGAGCACCTCTGGTAGAAAGACCCAAGGTCCGGGACCTACGGTCCCTG
- 7141 TCCCCGCAGGTTGCCCCGCCTGCCCCAGCGCTACTGGCAAATGCGGCCCCTGTTTCTGGA AGGGGCGTCCAACGGGGCGGACGGGGTCGCGATGACCGTTTACGCCGGGGACAAAGACCT

7167 ECO47III

- 7201 GCTGCTTGGGAACCACGCGCAGTGCCCCTACGGGGTGCTCCTCAAGACGCACTGCCCGCTCACGAGACCCTTGGTGCGCGTCACGGGGATGCCCCACGAGGAGTTCTGCGTGACGGGCGA
- 7261 GCGAGCTGCGGTCACCCCAGCAGCCGGTGTCTGTGCCCGGGAGAAGCCCCAGGGCTCTGT CGCTCGACGCCAGTGGGGTCGTCGGCCACAGACACGGGCCCTCTTCGGGGTCCCGAGACA
- 7321 GGCGGCCCCGAGGAGGAGACACAGACCCCCGTCGCCTGGTGCAGCTGCTCCGCCAGCA CCGCCGGGGGCTCCTCCTGTGTCTGGGGGCAGCGGACCACGTCGACGAGGCGGTCGT
- 7441 AGGCCTCTGGGGCTCCAGGCACAACGAACGCCGCTTCCTCAGGAACACCAAGAAGTTCAT TCCGGAGACCCCGAGGTCCGTGTTGCTTGCGGCGAAGGAGTCCTTGTGGTTCTTCAAGTA
- 7501 CTCCCTGGGGAAGCATGCCAAGCTCTCGCTGCAGGAGCTGACGTGGAAGATGAGCGTGCG GAGGGACCCCTTCGTACGGTTCGAGAGCGACGTCCTCGACTGCACCTTCTACTCGCACGC
- 7561 GGACTGCGCTGGCGCAGGAGCCCAGGTGAGGAGGTGGTGGCCGTCGAGGGCCCAGG CCTGACGCGAACCGACGCTCCTCGGGTCCACTCCTCCACCACCGGCAGCTCCCGGGTCC

7575 FSP1

Intron2

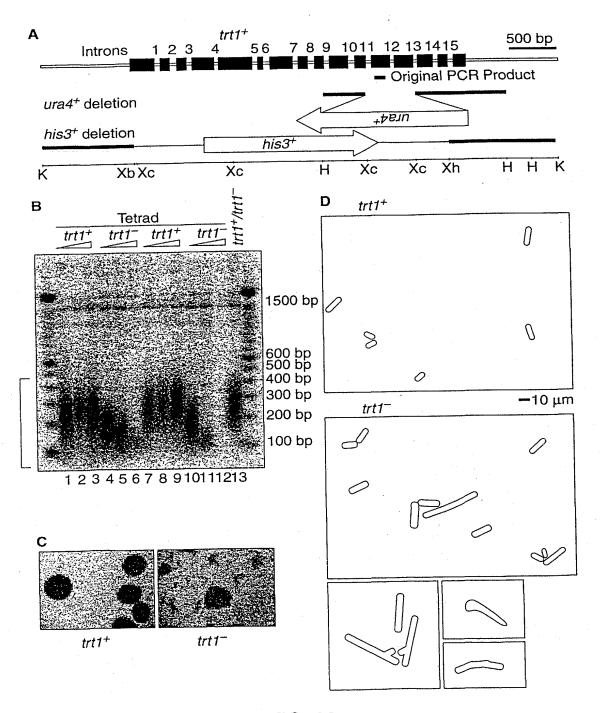
- 7681 GTCTCCATCGTCACGTGGGCACACGTGGCTTTTCGCTCAGGACGTCGAGTGGACACGGTGCACCGGAAAAGCGAGTCCTGCAGCTCACCTGTGCCAC

**>
7741 ATCGAGGTCGACTCTAGAGGATCCCCGGGTACCGAGCTCGAATTCGTAATCATGGTCATA
TAGCTCCAGCTGAGATCTCCTAGGGGCCCATGGCTCGAGCTTAAGCATTAGTACCAGTAT

7747 SAL1

FIG. 21 (CONTINUED)

32/34



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FIG. 22

Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
LOMERASE CATALYTIC SUBUNIT: DIAGNOSTIC
THERAPEUTIC METHODS
Sheet 33 of 34

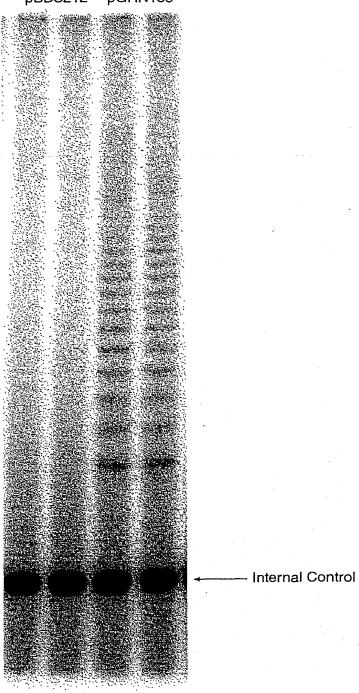
33/34

FIG. 23

TCTACCTTGACAGACCTCCAGCCGTACATGCGACAGTTCGTGGCTCACCTGCAGGAG ACCAGCCCGCTGAGGGATGCCGTCGTCATCGAGCAGAGCTCCTCCCTGAATGAGGCC AGCAGTGGCCTCTTCGACGTCTTCCTACGCTTCATGTGCCACCACGCCGTGCGCATC AGGGGCAAGTC

Atty. Docket No.: 015389-002640US
Applicant: Thomas R. Cech et al.
MERASE CATALYTIC SUBUNIT: DIAGNOSTIC A
THERAPEUTIC METHODS
Sheet 34 of 34

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